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a 58 amino acid amino-terminal insert and four microtubule binding domains). Table 1 summarizes the features of the six human tau isoforms, which are designated "A" through "F." The isoforms may also be respectively referred to as numbers "1" through "6," or by their total length in amino acid residues.

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### **IN THE SEQUENCE LISTING**

Please replace the sequence listing with the enclosed ~~pages~~ 1 – 14 containing the substitute Sequence Listing.

### **REMARKS**

The specification has been amended to insert a SEQ ID NO for the peptide appearing at page 5, paragraph [0023].

A substitute Sequence Listing is provided, listing SEQ ID NOs:1 through 16. SEQ ID NOs:1 through 15 are those listed in the original Sequence Listing, filed with the application on January 22, 2001. The substitute Sequence Listing includes SEQ ID NO:16, identifying the peptide appearing at page 5, paragraph [0023] of the specification as filed, as described above. No new matter has been added.

A paper copy of the substitute Sequence Listing is attached hereto. Also provided is a diskette containing a computer readable form of the substitute Sequence Listing. The information recorded in the computer readable form is identical to the paper copy of the substitute Sequence Listing.

### **Conclusion**

Applicants respectfully submit that claims 1 - 17 are in condition for allowance. A notice of allowance is earnestly solicited. Attached hereto is a marked-up version of the

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changes made to the specification by the current amendment. The attached pages are captioned  
"Version with markings to show changes made."

Respectfully submitted,



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Date: *April 25, 2002*

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**VERSION WITH MARKINGS TO SHOW CHANGES MADE**

**In the Specification**

**Paragraph [0023] on page 5:**

Tau protein microtubule binding domains, which contain the core microtubule binding domain motif proline-glycine-glycine-glycine (PGGG) [SEQ ID NO:16], are designated R1, R2, R3, and R4, and are encoded by exons 9, 10, 11, and 12, respectively. Exon 10, which encodes amino acid residues 275 through 305, is alternatively utilized (present in three of the six isoforms), such that the R2 microtubule binding domain is present only in tau isoforms containing four repeats. Thus, the isoforms range in size from 352 amino acid residues (with no amino-terminal inserts and three microtubule binding domains) to 441 amino acid residues (with a 58 amino acid amino-terminal insert and four microtubule binding domains). Table 1 summarizes the features of the six human tau isoforms, which are designated "A" through "F." The isoforms may also be respectively referred to as numbers "1" through "6," or by their total length in amino acid residues.

**In the Sequence Listing**

Please insert pages 1 – 14 containing the Sequence Listing.